- 149. A genetically modified host cell comprising the isolated nucleic acid of claim 24, or the recombinant expression vector of claim 28.
- **150.** A library of nucleic acids comprising a plurality of members, wherein each member nucleic acid comprises a nucleotide sequence encoding a DNA-targeting RNA comprising:
 - a) a first segment comprising a nucleotide sequence that is complementary to a target sequence in the target DNA;
 - b) a second segment that interacts with a site-directed polypeptide; and
 - c) a stability control sequence,
 - wherein each member differs from other members of the library in the nucleotide sequence of the first segment.
- **151.** A library of claim **150**, wherein the nucleotide sequence encoding a DNA-targeting RNA is operably linked to a promoter.
 - 152. A kit comprising:
 - a) a DNA-targeting RNA, or a nucleic acid comprising a nucleotide sequence encoding the DNA-targeting RNA, wherein the DNA-targeting RNA comprises:

- i)) a first segment comprising a nucleotide sequence that is complementary to a target sequence in the target DNA;
- ii)) a second segment that interacts with a site-directed polypeptide; and
- iii) a stability control sequence; and
- b) a buffer.
- **153**. The kit of claim **152**, further comprising a variant Cas9 site-directed polypeptide that exhibits reduced endode-oxyribonuclease activity relative to wild-type Cas9.
- **154.** The kit of claim **152**, wherein the nucleic acid comprising a nucleotide sequence encoding the DNA-targeting RNA further comprises a nucleotide sequence encoding a variant Cas9 site-directed polypeptide that exhibits reduced endodeoxyribonuclease activity relative to wild-type Cas9.
- **155.** The kit of claim **152,** further comprising a nucleic acid comprising a nucleotide sequence encoding a variant Cas9 site-directed polypeptide that exhibits reduced endodeoxyribonuclease activity relative to wild-type Cas9.

* * * * *